## Perspective

## Relationship of household air pollution from solid fuel combustion with tuberculosis?

Household air pollution (HAP) from combustion of solid biomass fuel (such as dried-dung, wood and crop residue) for cooking and heating purposes, has emerged as an important risk factor in the pathogenesis, prevalence, morbidity and mortality from chronic respiratory diseases such as chronic bronchitis (CB), chronic obstructive pulmonary disease (COPD) and asthma<sup>1-5</sup>. Exposure to HAP also accounts for frequent occurrence of respiratory infections and lung function impairment particularly in children<sup>6-8</sup>. There are inconsistent data on association of HAP with respiratory infections in adults. In the recent past, there have been reports on the role of HAP as a risk factor for tuberculosis (TB)<sup>9,10</sup>. But the evidence has been sparse, and mostly from the small studies.

It is known that TB occurs more often among inmates of slums with poor living conditions<sup>11-13</sup>. The poor living conditions include the environmental pollution from use of solid fuels for combustion for cooking and heating, widely prevalent in these areas. A systematic review of 33 studies on tobacco smoke, indoor air pollution and TB in 2007, including five on biomass fuel combustion concluded that the exposures increased the risk of TB, but stronger evidence was needed before the final conclusions could be drawn<sup>14</sup>.

Of the 994 full text articles, only 10 possible articles were found by the authors searching for an association of indoor fuel combustion (IAP) and TB<sup>15</sup>. Only three of the six studies which met the inclusion criteria for the systematic review reported a significant risk of TB amongst the exposed subjects<sup>15</sup>. In another meta-analysis in 2012, an extensive review of literature revealed 10 studies which showed a pooled odds ratio (OR) of 1.55 (95% C.I. 1.11 - 2.18) for TB in indoor air pollution (IAP) exposed individuals<sup>16</sup>. There were eight case control studies of which three did not show any

significant association<sup>16-19</sup>. On the other hand, both the cross-sectional studies from India as well as five case control studies from India, Benin, Mexico and Nepal, respectively showed significant risk of exposure<sup>16,20-26</sup>. Some of the studies had classified variable odds ratio with different types of fuels used in the kitchen or with reference to present and past exposure<sup>25,26</sup>. No association was shown in a recent report on 355 TB cases from Directly Observed Therapy Centres and 804 controls from the neighbourhoods of patients from rural settings of Sidama zone in South Ethiopia<sup>27</sup>. The study, however, suffered because of the low statistical power as pointed out by the authors themselves. The neighbourhood controls which they selected were over-matched and inappropriate to measure this 'slight' association in the presence of similarities of non-risk factor exposures<sup>27</sup>.

Most of the studies available on the subject are small in size based on a case-control design. Such a design is subject to variations depending upon the methods used to assess the exposure. This was evident by the presence of significant heterogeneity in different studies. None of these studies had employed any objective parameter of pollutant exposure. Some of the studies report an attempt at sub-categorization of exposure. The Benin study reported a higher risk (OR > 1) for use of solid-fuels for cooking on univariate analysis<sup>24</sup>. On multivariate analysis using other significant variables as well, the association was no longer significant<sup>24</sup>. Similarly, in a Delhi study, the TB risk was significantly influenced by the location of the kitchen and the presence of respiratory symptoms while cooking, but the TB odds did not significantly differ with the type of fuel used for cooking<sup>28</sup>. An earlier study from India reported a high OR of 3.26 (95% CI 1.25 - 8.46) for TB with "not having a separate kitchen" while the authors had

**Box.** A partial list of chemical constituents common to the smoke from tobacco and biomass fuels

- Particulates (PM<sub>10</sub>, PM<sub>2.5</sub>)
  - Dibenzopyrenes
- · Carbon monoxide
- 1-3 Butadiene
- Ammonia
- Fenols
- Formaldehyde
- Cadmium
- Benzene
- Arsenic
- Benzo [a] pyrene

Source: Refs 33-35

also reported no significant association with the use of biomass fuels<sup>18</sup>. Such observations only point to the difficulties of exposure assessment. It is not uncommon in India and many other developing countries to have a working indoor kitchen to cook food within the one or two room living dwellings or huts. It is presumable that the environment of such dwellings gets highly polluted from combustion of solid (biomass) fuels.

There are several limitations to conduct large epidemiological studies because of the numbers of variables and other confounders which play an important role in causing TB. History of tobacco smoking, consumption of alcohol and presence of co-morbidities such as diabetes and human immunodeficiency virus (HIV) infection are some of the important risk factors for TB<sup>29,30</sup>. The chances of infection with mycobacteria are also high among contacts of TB patients living closely in crowded homes with poor ventilation. Overcrowding, urbanization and industrialization in highly populated cities and towns are also responsible for an increased incidence of TB<sup>31,32</sup>.

There are a large number of constituents of smoke emanating from combustion of solid, biomass fuels which are similar to those of tobacco smoke. A partial list of some toxic pollutants of tobacco smoke selected from the Federal Register, which are common to those of solid fuels taken from other sources is provided<sup>33-35</sup>. This points to the obvious possibility of common toxicities such as carcinogenesis, increased allergic sensitization and reduced pulmonary defenses. Besides these constituents, there are several other noxious constituents of solid fuels such as gaseous products (ozone, nitric oxide, nitrogen dioxide, carbon dioxide, sulphur oxides) and metals (lead, manganese, selenium, calcium, mercury, silicon) which are likely to produce damage to the respiratory tract and impair mucociliary defenses34,35.

The concentrations of the pollutants in the indoor air are particularly high as has been shown in numerous studies from several developing countries<sup>34-36</sup>. The duration of exposure to the indoor air is prolonged and persistent. These factors make the lungs continuously exposed to inhaled air, as more vulnerable to the harmful effects of pollutants, such as the lower respiratory tract infections, including tuberculosis. There are experimental data which show the impairment of immune defenses on exposure to wood smoke. Inhalation of wood smoke in rabbits caused impairment of macrophage function, impaired mucociliary function and reduced bacterial clearance<sup>37-39</sup>. All these functions are known to increase the chances of TB infection.

There are no studies on the effects of exposure to biomass fuel combustion on clinical features and mortality from tuberculosis. Similarly, influence of exposure on treatment effects and outcomes are not known. Assumptions can however, be made based on the known harmful effects of the combustion-smoke on the respiratory symptoms. Exposure to biomass combustion smoke is independently responsible for the presence of increased respiratory symptoms, chronic bronchitis and asthma<sup>4,5,40,41</sup>. TB patients, therefore, in the presence of continued exposure to indoor combustion of biomass fuels are likely to suffer from exaggerated and prolonged symptoms and disease complications. Tobacco smokers with TB are more likely to suffer from chronic cough and sputum smear positive tuberculosis<sup>42</sup>. Tobacco smoking has been comincingly shown to relate to TB mortality<sup>43-47</sup>. If one takes the smoking analogy, it can be presumed that the outcomes are likely to be poorer and mortality higher in the IAP exposed patients.

In conclusion, the indoor exposure to biomass fuel combustion used for heating and cooking is an identifiable risk factor for TB with demonstrable higher odds ratios in most studies. This is a subject which needs further investigations to clearly define the disease association, morbidity and mortality. Because of the potential implications as a treatment modifying factor and on outcomes of therapy,the issue has also got an important role in TB control programmes from public health point of view.

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